

## **RAW SEQUENCE LISTING**

**The Biotechnology Systems Branch of the Scientific and Technical Information Center (STIC) no errors detected.**

Application Serial Number: 10/712,795A  
Source: IFW/6  
Date Processed by STIC: 7/19/06

# ***ENTERED***



IFW16

## RAW SEQUENCE LISTING

DATE: 07/19/2006

PATENT APPLICATION: US/10/712,795A

TIME: 09:01:20

Input Set : A:\DOC0216USSEQ2.txt

Output Set: N:\CRF4\07192006\J712795A.raw

3 <110> APPLICANT: Rosanne M. Crooke  
 4 Mark J. Graham  
 5 Kristina Lemonidis Tarbet  
 6 Kenneth W. Dobie  
 7 Susan M. Freier  
 9 <120> TITLE OF INVENTION: ANTISENSE MODULATION OF APOLIPOPROTEIN B EXPRESSION  
 12 <130> FILE REFERENCE: DOC-0216US(ISIS.003CP1)  
 14 <140> CURRENT APPLICATION NUMBER: US 10/712,795A  
 15 <141> CURRENT FILING DATE: 2003-11-13  
 17 <150> PRIOR APPLICATION NUMBER: US 60/426,234  
 18 <151> PRIOR FILING DATE: 2002-11-13  
 20 <150> PRIOR APPLICATION NUMBER: PCT/US03/15493  
 21 <151> PRIOR FILING DATE: 2003-05-15  
 23 <160> NUMBER OF SEQ ID NOS: 897  
 25 <210> SEQ ID NO: 1  
 26 <211> LENGTH: 20  
 27 <212> TYPE: DNA  
 28 <213> ORGANISM: Artificial Sequence  
 30 <220> FEATURE:  
 31 <223> OTHER INFORMATION: Antisense Oligonucleotide  
 33 <400> SEQUENCE: 1  
 34 tccgtcatcg ctcttcaggg 20  
 36 <210> SEQ ID NO: 2  
 37 <211> LENGTH: 20  
 38 <212> TYPE: DNA  
 39 <213> ORGANISM: Artificial Sequence  
 41 <220> FEATURE:  
 42 <223> OTHER INFORMATION: Antisense Oligonucleotide  
 44 <400> SEQUENCE: 2  
 45 atgcattctg cccccaagga 20  
 47 <210> SEQ ID NO: 3  
 48 <211> LENGTH: 14121  
 49 <212> TYPE: DNA  
 50 <213> ORGANISM: Homo sapiens  
 52 <220> FEATURE:  
 53 <221> NAME/KEY: CDS  
 54 <222> LOCATION: (129)..(13820)  
 56 <400> SEQUENCE: 3  
 57 attcccaccg ggacctgcgg ggctgagtgc ctttctcggt tgctgccgct gaggagcccg 60  
 59 cccagccagc cagggccgcg aggccgaggc cagggccgcag cccaggagcc gccccaccgc 120  
 61 agctggcg atg gac ccg ccg agg ccc gcg ctg ctg gcg ctg gcg ctg 170  
 62 Met Asp Pro Pro Arg Pro Ala Leu Leu Ala Leu  
 63 1 5 10

## RAW SEQUENCE LISTING

PATENT APPLICATION: US/10/712,795A

DATE: 07/19/2006

TIME: 09:01:20

Input Set : A:\DOC0216USSEQ2.txt

Output Set : N:\CRF4\07192006\J712795A.raw

```

65 cct gcg ctg ctg ctg ctg ctg gcg ggc gcc agg gcc gaa gag gaa 218
66 Pro Ala Leu Leu Leu Leu Leu Leu Ala Gly Ala Arg Ala Glu Glu Glu
67 15 20 25 30
69 atg ctg gaa aat gtc agc ctg gtc tgt cca aaa gat gcg acc cga ttc 266
70 Met Leu Glu Asn Val Ser Leu Val Cys Pro Lys Asp Ala Thr Arg Phe
71 35 40 45
74 aag cac ctc cgg aag tac aca tac aac tat gag gct gag agt tcc agt 314
75 Lys His Leu Arg Lys Tyr Thr Tyr Asn Tyr Glu Ala Glu Ser Ser Ser
76 50 55 60
78 gga gtc cct ggg act gct gat tca aga agt gcc acc agg atc aac tgc 362
79 Gly Val Pro Gly Thr Ala Asp Ser Arg Ser Ala Thr Arg Ile Asn Cys
80 65 70 75
82 aag gtt gag ctg gag gtt ccc cag ctg tgc agc ttc atc ctg aag acc 410
83 Lys Val Glu Leu Glu Val Pro Gln Leu Cys Ser Phe Ile Leu Lys Thr
84 80 85 90
86 agc cag tgc acc ctg aaa gag gtg tat ggc ttc aac cct gag ggc aaa 458
87 Ser Gln Cys Thr Leu Lys Glu Val Tyr Gly Phe Asn Pro Glu Gly Lys
88 95 100 105 110
90 gcc ttg ctg aag aaa acc aag aac tct gag gag ttt gct gca gcc atg 506
91 Ala Leu Leu Lys Lys Thr Lys Asn Ser Glu Glu Phe Ala Ala Ala Met
92 115 120 125
94 tcc agg tat gag ctg aag ctg gcc att cca gaa ggg aag cag gtt ttc 554
95 Ser Arg Tyr Glu Leu Lys Leu Ala Ile Pro Glu Gly Lys Gln Val Phe
96 130 135 140
98 ctt tac ccg gag aaa gat gaa cct act tac atc ctg aac atc aag agg 602
99 Leu Tyr Pro Glu Lys Asp Glu Pro Thr Tyr Ile Leu Asn Ile Lys Arg
100 145 150 155
102 ggc atc att tct gcc ctg ctg gtt ccc cca gag aca gaa gaa gcc aag 650
103 Gly Ile Ser Ala Leu Leu Val Pro Pro Glu Thr Glu Glu Ala Lys
104 160 165 170
106 caa gtg ttg ttt ctg gat acc gtg tat gga aac tgc tcc act cac ttt 698
107 Gln Val Leu Phe Leu Asp Thr Val Tyr Gly Asn Cys Ser Thr His Phe
108 175 180 185 190
110 acc gtc aag acg agg aag ggc aat gtg gca aca gaa ata tcc act gaa 746
111 Thr Val Lys Thr Arg Lys Gly Asn Val Ala Thr Glu Ile Ser Thr Glu
112 195 200 205
114 aga gac ctg ggg cag tgt gat cgc ttc aag ccc atc cgc aca ggc atc 794
115 Arg Asp Leu Gly Gln Cys Asp Arg Phe Lys Pro Ile Arg Thr Gly Ile
116 210 215 220
118 agc cca ctt gct ctg atc aaa ggc atg acc cgc ccc ttg tca act ctg 842
119 Ser Pro Leu Ala Leu Ile Lys Gly Met Thr Arg Pro Leu Ser Thr Leu
120 225 230 235
122 atc agc agc agc cag tcc tgt cag tac aca ctg gac gct aag agg aag 890
123 Ile Ser Ser Ser Gln Ser Cys Gln Tyr Thr Leu Asp Ala Lys Arg Lys
124 240 245 250
126 cat gtg gca gaa gcc atc tgc aag gag caa cac ctg ttc ctg cct ttc 938
127 His Val Ala Glu Ala Ile Cys Lys Glu Gln His Leu Phe Leu Pro Phe
128 255 260 265 270
130 tcc tac aac aat aag tat ggg atg gta gca caa gtg aca cag act ttg 986

```

## RAW SEQUENCE LISTING

DATE: 07/19/2006

PATENT APPLICATION: US/10/712,795A

TIME: 09:01:20

Input Set : A:\DOC0216USSEQ2.txt

Output Set: N:\CRF4\07192006\J712795A.raw

131	Ser	Tyr	Asn	Asn	Lys	Tyr	Gly	Met	Val	Ala	Gln	Val	Thr	Gln	Thr	Leu	
132					275					280					285		
134	aaa	ctt	gaa	gac	aca	cca	aag	atc	aac	agc	cgc	ttc	ttt	ggg	gaa	ggg	1034
135	Lys	Leu	Glu	Asp	Thr	Pro	Lys	Ile	Asn	Ser	Arg	Phe	Phe	Gly	Glu	Gly	
136				290					295					300			
138	act	aag	aag	atg	ggc	ctc	gca	ttt	gag	agc	acc	aaa	tcc	aca	tca	cct	1082
139	Thr	Lys	Lys	Met	Gly	Leu	Ala	Phe	Glu	Ser	Thr	Lys	Ser	Thr	Ser	Pro	
140			305					310					315				
142	cca	aag	cag	gcc	gaa	gct	gtt	ttg	aag	act	ctc	cag	gaa	ctg	aaa	aaa	1130
143	Pro	Lys	Gln	Ala	Glu	Ala	Val	Leu	Lys	Thr	Leu	Gln	Glu	Leu	Lys	Lys	
144		320					325					330					
146	cta	acc	atc	tct	gag	caa	aat	atc	cag	aga	gct	aat	ctc	ttc	aat	aag	1178
147	Leu	Thr	Ile	Ser	Glu	Gln	Asn	Ile	Gln	Arg	Ala	Asn	Leu	Phe	Asn	Lys	
148	335				340					345					350		
150	ctg	gtt	act	gag	ctg	aga	ggc	ctc	agt	gat	gaa	gca	gtc	aca	tct	ctc	1226
151	Leu	Val	Thr	Glu	Leu	Arg	Gly	Leu	Ser	Asp	Glu	Ala	Val	Thr	Ser	Leu	
152				355					360					365			
154	ttg	cca	cag	ctg	att	gag	gtg	tcc	agc	ccc	atc	act	tta	caa	gcc	ttg	1274
155	Leu	Pro	Gln	Leu	Ile	Glu	Val	Ser	Ser	Pro	Ile	Thr	Leu	Gln	Ala	Leu	
156			370					375					380				
158	gtt	cag	tgt	gga	cag	cct	cag	tgc	tcc	act	cac	atc	ctc	cag	tgg	ctg	1322
159	Val	Gln	Cys	Gly	Gln	Pro	Gln	Cys	Ser	Thr	His	Ile	Leu	Gln	Trp	Leu	
160			385				390					395					
162	aaa	cgt	gtg	cat	gcc	aac	ccc	ctt	ctg	ata	gat	gtg	gtc	acc	tac	ctg	1370
163	Lys	Arg	Val	His	Ala	Asn	Pro	Leu	Leu	Ile	Asp	Val	Val	Thr	Tyr	Leu	
164		400				405					410						
166	gtg	gcc	ctg	atc	ccc	gag	ccc	tca	gca	cag	cag	ctg	cga	gag	atc	ttc	1418
167	Val	Ala	Leu	Ile	Pro	Glu	Pro	Ser	Ala	Gln	Gln	Leu	Arg	Glu	Ile	Phe	
168	415				420					425				430			
170	aac	atg	gcg	agg	gat	cag	cgc	agc	cga	gcc	acc	ttg	tat	gcg	ctg	agc	1466
171	Asn	Met	Ala	Arg	Asp	Gln	Arg	Ser	Arg	Ala	Thr	Leu	Tyr	Ala	Leu	Ser	
172				435				440					445				
174	cac	gcg	gtc	aac	aac	tat	cat	aag	aca	aac	cct	aca	ggg	acc	cag	gag	1514
175	His	Ala	Val	Asn	Asn	Tyr	His	Lys	Thr	Asn	Pro	Thr	Gly	Thr	Gln	Glu	
176			450					455					460				
178	ctg	ctg	gac	att	gct	aat	tac	ctg	atg	gaa	cag	att	caa	gat	gac	tgc	1562
179	Leu	Leu	Asp	Ile	Ala	Asn	Tyr	Leu	Met	Glu	Gln	Ile	Gln	Asp	Asp	Cys	
180			465				470					475					
182	act	ggg	gat	gaa	gat	tac	acc	tat	ttg	att	ctg	cgg	gtc	att	gga	aat	1610
183	Thr	Gly	Asp	Glu	Asp	Tyr	Thr	Tyr	Leu	Ile	Leu	Arg	Val	Ile	Gly	Asn	
184		480				485					490						
186	atg	ggc	caa	acc	atg	gag	cag	tta	act	cca	gaa	ctc	aag	tct	tca	atc	1658
187	Met	Gly	Gln	Thr	Met	Glu	Gln	Leu	Thr	Pro	Glu	Leu	Lys	Ser	Ser	Ile	
188	495				500					505				510			
190	ctc	aaa	tgt	gtc	caa	agt	aca	aag	cca	tca	ctg	atg	atc	cag	aaa	gct	1706
191	Leu	Lys	Cys	Val	Gln	Ser	Thr	Lys	Pro	Ser	Leu	Met	Ile	Gln	Lys	Ala	
192				515				520					525				
194	gcc	atc	cag	gct	ctg	cgg	aaa	atg	gag	cct	aaa	gac	aag	gac	cag	gag	1754
195	Ala	Ile	Gln	Ala	Leu	Arg	Lys	Met	Glu	Pro	Lys	Asp	Lys	Asp	Gln	Glu	

## RAW SEQUENCE LISTING

DATE: 07/19/2006

PATENT APPLICATION: US/10/712,795A

TIME: 09:01:20

Input Set : A:\DOC0216USSEQ2.txt

Output Set: N:\CRF4\07192006\J712795A.raw

196		530		535		540		
198	gtt	ctt	ctt	cag	act	ttc	ctt	gat
199	Val	Leu	Leu	Gln	Thr	Phe	Leu	Asp
200		545		550		555		
202	ctg	gct	gcc	tat	ctt	atg	ttg	atg
203	Leu	Ala	Ala	Tyr	Leu	Met	Leu	Met
204		560		565		570		
206	aac	aaa	att	gtc	caa	att	cta	cca
207	Asn	Lys	Ile	Val	Gln	Ile	Leu	Pro
208	575			580		585		590
210	aac	ttt	gtg	gct	tcc	cat	att	gcc
211	Asn	Phe	Val	Ala	Ser	His	Ile	Ala
212				595		600		605
214	gat	atc	caa	gat	ctg	aaa	aag	tta
215	Asp	Ile	Gln	Asp	Leu	Lys	Lys	Leu
216			610		615		620	
218	caa	ctt	cca	act	gtc	atg	gac	ttc
219	Gln	Leu	Pro	Thr	Val	Met	Asp	Phe
220		625		630		635		
222	ctc	tac	aaa	tct	gtt	tct	ctt	cca
223	Leu	Tyr	Lys	Ser	Val	Ser	Leu	Pro
224		640		645		650		
226	ata	gaa	ggg	aat	ctt	ata	ttt	gat
227	Ile	Glu	Gly	Asn	Leu	Ile	Phe	Asp
228	655			660		665		670
231	agc	atg	ctg	aaa	act	acc	ctc	act
232	Ser	Met	Leu	Lys	Thr	Thr	Leu	Thr
233				675		680		685
235	ctc	atc	gag	att	ggc	ttg	gaa	gga
236	Leu	Ile	Glu	Ile	Gly	Leu	Glu	Gly
237			690		695		700	
239	gct	ctt	ttt	ggg	aag	caa	gga	ttt
240	Ala	Leu	Phe	Gly	Lys	Gln	Gly	Phe
241		705		710		715		
243	ttg	tac	tgg	gtt	aat	ggt	caa	gtt
244	Leu	Tyr	Trp	Val	Asn	Gly	Gln	Val
245		720		725		730		
247	gtg	gac	cac	ttt	ggc	tat	acc	aaa
248	Val	Asp	His	Phe	Gly	Tyr	Thr	Lys
249	735			740		745		750
251	gta	aat	gga	ata	atg	ctc	agt	gtt
252	Val	Asn	Gly	Ile	Met	Leu	Ser	Val
253			755		760		765	
255	tcc	aaa	gaa	gtc	ccg	gaa	gcc	aga
256	Ser	Lys	Glu	Val	Pro	Glu	Ala	Arg
257			770		775		780	
259	gag	ctt	ggt	ttt	gcc	agt	ctc	cat
260	Glu	Leu	Gly	Phe	Ala	Ser	Leu	His
261		785		790		795		

## RAW SEQUENCE LISTING

PATENT APPLICATION: US/10/712,795A

DATE: 07/19/2006

TIME: 09:01:20

Input Set : A:\DOC0216USSEQ2.txt

Output Set: N:\CRF4\07192006\J712795A.raw

263	ctt	ctg	atg	ggt	gcc	cgc	act	ctg	cag	ggg	atc	ccc	cag	atg	att	gga	2570
264	Leu	Leu	Met	Gly	Ala	Arg	Thr	Leu	Gln	Gly	Ile	Pro	Gln	Met	Ile	Gly	
265		800					805					810					
267	gag	gtc	atc	agg	aag	ggc	tca	aag	aat	gac	ttt	ttt	ctt	cac	tac	atc	2618
268	Glu	Val	Ile	Arg	Lys	Gly	Ser	Lys	Asn	Asp	Phe	Phe	Leu	His	Tyr	Ile	
269	815					820					825					830	
271	ttc	atg	gag	aat	gcc	ttt	gaa	ctc	ccc	act	gga	gct	gga	tta	cag	ttg	2666
272	Phe	Met	Glu	Asn	Ala	Phe	Glu	Leu	Pro	Thr	Gly	Ala	Gly	Leu	Gln	Leu	
273					835					840					845		
275	caa	ata	tct	tca	tct	gga	gtc	att	gct	ccc	gga	gcc	aag	gct	gga	gta	2714
276	Gln	Ile	Ser	Ser	Ser	Gly	Val	Ile	Ala	Pro	Gly	Ala	Lys	Ala	Gly	Val	
277					850					855					860		
279	aaa	ctg	gaa	gta	gcc	aac	atg	cag	gct	gaa	ctg	gtg	gca	aaa	ccc	tcc	2762
280	Lys	Leu	Glu	Val	Ala	Asn	Met	Gln	Ala	Glu	Leu	Val	Ala	Lys	Pro	Ser	
281			865					870						875			
283	gtg	tct	gtg	gag	ttt	gtg	aca	aat	atg	ggc	atc	atc	att	ccg	gac	ttc	2810
284	Val	Ser	Val	Glu	Phe	Val	Thr	Asn	Met	Gly	Ile	Ile	Ile	Pro	Asp	Phe	
285		880					885							890			
287	gct	agg	agt	ggg	gtc	cag	atg	aac	acc	aac	ttc	ttc	cac	gag	tcg	ggt	2858
288	Ala	Arg	Ser	Gly	Val	Gln	Met	Asn	Thr	Asn	Phe	Phe	His	Glu	Ser	Gly	
289	895					900					905					910	
291	ctg	gag	gct	cat	gtt	gcc	cta	aaa	gct	ggg	aag	ctg	aag	ttt	atc	att	2906
292	Leu	Glu	Ala	His	Val	Ala	Leu	Lys	Ala	Gly	Lys	Leu	Lys	Phe	Ile	Ile	
293					915					920					925		
295	cct	tcc	cca	aag	aga	cca	gtc	aag	ctg	ctc	agt	gga	ggc	aac	aca	tta	2954
296	Pro	Ser	Pro	Lys	Arg	Pro	Val	Lys	Leu	Leu	Ser	Gly	Gly	Asn	Thr	Leu	
297					930					935					940		
299	cat	ttg	gtc	tct	acc	acc	aaa	acg	gag	gtg	atc	cca	cct	ctc	att	gag	3002
300	His	Leu	Val	Ser	Thr	Thr	Lys	Thr	Glu	Val	Ile	Pro	Pro	Leu	Ile	Glu	
301			945					950						955			
303	aac	agg	cag	tcc	tgg	tca	gtt	tgc	aag	caa	gtc	ttt	cct	ggc	ctg	aat	3050
304	Asn	Arg	Gln	Ser	Trp	Ser	Val	Cys	Lys	Gln	Val	Phe	Pro	Gly	Leu	Asn	
305		960					965							970			
307	tac	tgc	acc	tca	ggc	gct	tac	tcc	aac	gcc	agc	tcc	aca	gac	tcc	gcc	3098
308	Tyr	Cys	Thr	Ser	Gly	Ala	Tyr	Ser	Asn	Ala	Ser	Ser	Thr	Asp	Ser	Ala	
309	975					980					985					990	
311	tcc	tac	tat	ccg	ctg	acc	ggg	gac	acc	aga	tta	gag	ctg	gaa	ctg	agg	3146
312	Ser	Tyr	Tyr	Pro	Leu	Thr	Gly	Asp	Thr	Arg	Leu	Glu	Leu	Glu	Leu	Arg	
313					995					1000					1005		
315	cct	aca	gga	gag	att	gag	cag	tat	tct	gtc	agc	gca	acc	tat	gag	ctc	3194
316	Pro	Thr	Gly	Glu	Ile	Glu	Gln	Tyr	Ser	Val	Ser	Ala	Thr	Tyr	Glu	Leu	
317					1010					1015					1020		
319	cag	aga	gag	gac	aga	gcc	ttg	gtg	gat	acc	ctg	aag	ttt	gta	act	caa	3242
320	Gln	Arg	Glu	Asp	Arg	Ala	Leu	Val	Asp	Thr	Leu	Lys	Phe	Val	Thr	Gln	
321			1025					1030						1035			
323	gca	gaa	ggt	gcg	aag	cag	act	gag	gct	acc	atg	aca	ttc	aaa	tat	aat	3290
324	Ala	Glu	Gly	Ala	Lys	Gln	Thr	Glu	Ala	Thr	Met	Thr	Phe	Lys	Tyr	Asn	
325		1040					1045							1050			
327	cgg	cag	agt	atg	acc	ttg	tcc	agt	gaa	gtc	caa	att	ccg	gat	ttt	gat	3338

RAW SEQUENCE LISTING ERROR SUMMARY      DATE: 07/19/2006  
PATENT APPLICATION: US/10/712,795A      TIME: 09:01:21

Input Set : A:\DOC0216USSEQ2.txt  
Output Set: N:\CRF4\07192006\J712795A.raw

Please Note:

Use of n and/or Xaa have been detected in the Sequence Listing. Please review the Sequence Listing to ensure that a corresponding explanation is presented in the <220> to <223> fields of each sequence which presents at least one n or Xaa.

Seq#:503; N Pos. 44,99,156,468

Seq#:810; N Pos. 45,118,148,173,180

Seq#:855; N Pos. 7,469,470,471,472,473,474,476,477,478,479

Seq#:890; N Pos. 9

**VERIFICATION SUMMARY**

PATENT APPLICATION: US/10/712,795A

DATE: 07/19/2006

TIME: 09:01:21

Input Set : A:\DOC0216USSEQ2.txt

Output Set: N:\CRF4\07192006\J712795A.raw

L:9405 M:341 W: (46) "n" or "Xaa" used, for SEQ ID#:503 after pos.:0  
M:341 Repeated in SeqNo=503  
L:11951 M:341 W: (46) "n" or "Xaa" used, for SEQ ID#:810 after pos.:0  
M:341 Repeated in SeqNo=810  
L:12513 M:341 W: (46) "n" or "Xaa" used, for SEQ ID#:855 after pos.:0  
M:341 Repeated in SeqNo=855  
L:13143 M:341 W: (46) "n" or "Xaa" used, for SEQ ID#:890 after pos.:0